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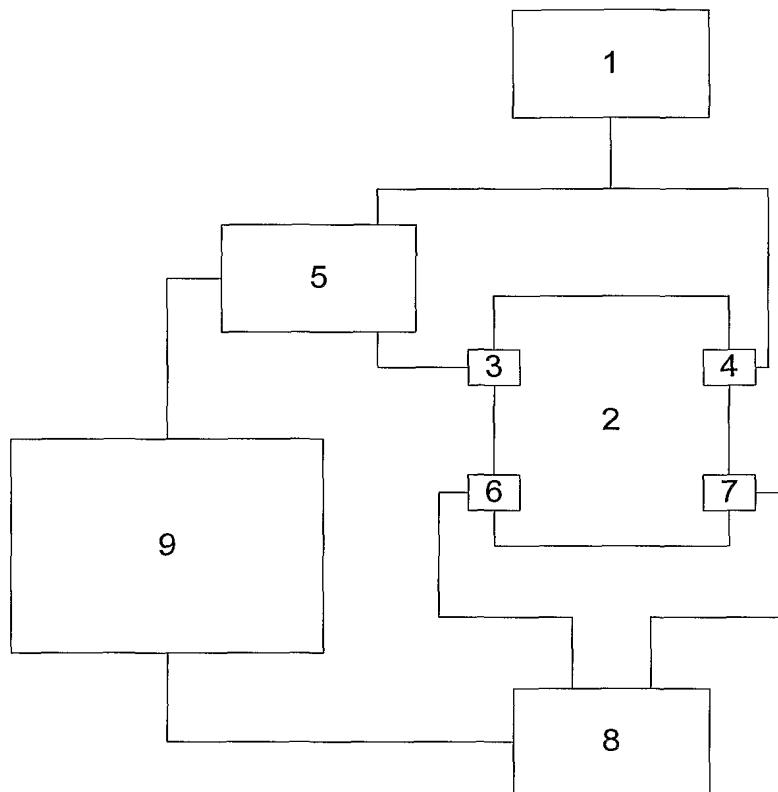
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(54) Title: MULTIFREQUENCY BIOIMPEDANCE DETERMINATION



(57) Abstract: The present invention provides a method of determining the impedance of a subject. This involves applying an electrical signal representing a range of superposed frequencies, and then determining the current flow through and voltage across the subject for a number of the frequencies within the range. The impedance of the subject is then determined at each of the number of frequencies.

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MULTIFREQUENCY BIOIMPEDANCE DETERMINATION

Background of the Invention

The present invention relates to a method and apparatus for determining the impedance of
5 a subject, and in particular to determining the biological impedance of a biological subject.

Description of the Prior Art

The reference to any prior art in this specification is not, and should not be taken as, an
acknowledgement or any form of suggestion that the prior art forms part of the common
10 general knowledge.

Correlations between whole-body impedance measurements and various body
characteristics, such as total body water (TBW) and fat-free mass (FFM), are
experimentally well established. As a consequence, bioelectrical impedance analysis
15 (BIA) is widely used in human nutrition and clinical research.

It is generally accepted that BIA provides a reliable estimate of total body water under
most conditions and in the National Institutes of Health Technology Assessment Statement
entitled "Bioelectrical Impedance Analysis in Body Composition Measurement, December
20 12-14, 1994" it was noted that BIA can be a useful technique for body composition
analysis in healthy individuals and in those with a number of chronic conditions such as
mild-to-moderate obesity, diabetes mellitus, and other medical conditions in which major
disturbances of water distribution are not prominent. In addition, BIA is fast, inexpensive,
and does not require extensive operator training or cross-validation.

25

BIA measures the impedance or opposition to the flow of an electric current through the
body fluids contained mainly in the lean and fat tissue. Impedance is low in lean tissue,
where intracellular fluid and electrolytes are primarily contained, but high in fat tissue.
Impedance is thus proportional to TBW.

30

Currently, in practice, a small constant current, typically 800 μ A at a fixed frequency,
usually 50 kHz, is passed between electrodes spanning the body and the voltage drop
between electrodes provides a measure of impedance. Prediction equations, previously

generated by correlating impedance measures against an independent estimate of TBW, may be used subsequently to convert a measured impedance to a corresponding estimate of TBW. Lean body mass is then calculated from this estimate using an assumed hydration fraction for lean tissue. Fat mass is calculated as the difference between body weight and
5 lean body mass.

The impedance of a biological tissue comprises two components, resistance and reactance. The conductive characteristics of body fluids provide the resistive component, whereas the cell membranes, acting as imperfect capacitors, contribute a frequency-dependent reactive
10 component. Impedance measurements made over a range of low to high (1 MHz) frequencies therefore allow development of prediction equations relating impedance measures at low frequencies to extracellular fluid volume and at high frequencies to total body fluid volume. This is often referred to as multi-frequency bioelectrical impedance analysis.
15

Recent applications of BIA increasingly use multi-frequency measurements, or a frequency spectrum, to evaluate differences in body composition caused by clinical and nutritional status. While the National Institutes of Health Technology Assessment Statement did not support the use of BIA under conditions that alter the normal relationship between the
20 extracellular (ECW) and intracellular water (ICW) compartments, recent studies indicate that the only model that accurately predicted change in ECW, ICW, and TBW is the zero-infinity kHz parallel multiple frequency model, often referred to as a Cole-Cole plot (example, refer Gudivaka, R., D.A. Schoeller, R. F. Kushner, and M. J. G. Bolt. Single- and multi-frequency models for bioelectrical impedance analysis of body water
25 compartments. *J. Appl.Physiol.* 87(3): 1087–1096, 1999).

Currently techniques for implementing multi-frequency analysis involve applying a number of signals to the subject in turn, with each signal having a respective frequency. The resulting impedance at each frequency is then determined separately, allowing the
30 dependence of impedance on frequency to be determined. An example of apparatus suitable for performing impedance determination using this technique is shown in US-5,280,429.

In this case, once the impedance at each frequency has been obtained, and the results are plotted as a graph of resistance versus frequency, reactance versus frequency, or resistance versus reactance (the zero-infinity kHz parallel multiple frequency plot, or Cole-Cole plot, referred to above).

5

However, this technique suffers from a number of drawbacks. In particular, it is necessary to generate a large number of data points for accurate plots to be made. Furthermore, as each respective frequency signal must be applied to the subject in turn, this procedure can take a long time, and in particular, can take as long as half-an-hour.

10

Summary of the Present Invention

In a first broad form the present invention provides a method of determining the impedance of a subject, the method including:

- a) Applying an electrical signal representing a range of superposed frequencies;
- 15 b) Determining for a number of frequencies within the range:
 - i) The current flow through the subject; and,
 - ii) The voltage across the subject; and,
- c) Determining the impedance of the subject at each of the number of frequencies.

20 The method typically includes:

- a) Generating component signals, each component signal having a respective one of the number of frequencies; and,
- b) Superposing the component signals to generate the electrical signal.

25 However, alternatively the electrical signal can be formed from white noise. In this case, the method typically includes:

- a) Generating the white noise using a Linear Feedback Shift Register (LFSR) circuit to produce a pseudo-random digital sequence; and,
- b) Converting the pseudo-random digital sequence to an analogue signal using a

30 digital to analogue (D/A) converter; and,

- c) Applying the analogue signal to the subject.

The method of determining the current flow generally includes:

- a) Sampling the current of the electrical signal applied to the subject; and,
- b) Converting the current signal to a digitised current signal.

The method of determining the voltage generally includes:

- 5 a) Obtaining a signal representing the voltage generated across the subject;
- b) Converting the voltage signal to a digitised voltage signal.

The method can include digitising the current and voltage signals by sampling the signals at a predetermined rate. Furthermore, the method can include digitising the current and
10 voltage signals by sampling the signals with a predetermined sample length. It will be appreciated that a range of values may be used for the predetermined rate, such as several MHz, with the sample length typically being up to a thousand or so sample points or more, depending on the implementation.

15 The method typically includes converting each of the digitised voltage and current signals into the frequency domain. This conversion may be performed using a Fast Fourier Transform (FFT).

The method can include using a processing system to:

- 20 a) Receive the converted voltage and current signals; and,
- b) Determine the impedance of the subject at each of the number of frequencies.

The processing system can be further adapted to determine the variation in the impedance with the frequency of the applied signal.

25

The method typically further includes generating a graphical representation of the variation in the impedance with the frequency of the applied signal.

30 In a second broad form the present invention provides apparatus for determining the impedance of a subject, the apparatus including:

- a) A signal generator for applying an electrical signal representing a range of superposed frequencies;

b) A voltage detector for determining the voltage across the subject at a number of frequencies within the range;

c) A current detector for determining the current flow through the subject at a number of frequencies within the range; and,

5 d) A processing system for determining the impedance of the subject at each of the number of frequencies.

In this case, the signal generator can be adapted to:

10 a) Generate component signals, each component signal having a respective one of the number of frequencies; and,

b) Superpose the component signals to generate the electrical signal.

Alternatively, the electrical signal can be formed from white noise, in which case the signal generator typically includes:

15 a) A shift register circuit to produce a pseudo-random digital sequence; and,

b) A D/A converter for converting the pseudo-random digital sequence to an analogue signal.

The shift register circuit typically includes:

20 a) A shift register having an output coupled to the D/A converter; and,

b) An exclusive OR (XOR) gate adapted to:

- i) Receive inputs from a number of predetermined locations in the first register;
- ii) Logically combine the inputs to generate an XOR output; and,
- iii) Provide the XOR output to an input of the shift register;

25

The signal generator can include a second shift register, the second shift register being adapted to couple an output of the first shift register to an input of the D/A converter.

The current detector typically includes:

30 a) A current sampler coupled to the signal generator for sampling the current flowing through the subject; and,

b) A current analogue to digital (A/D) converter for converting the sampled current to a digitised current signal.

The voltage detector generally includes a voltage A/D converter coupled to the subject via a respective set of electrodes, the voltage A/D converter being adapted to generate a digitised voltage signal.

5

The current and voltage A/D converters may be adapted to digitise the current and voltage signals by sampling the signals at a predetermined rate, and/or by sampling the signals with a predetermined sample length. As mentioned above however, alternative sample rates and lengths may be used.

10

The processing system is preferably adapted to convert each of the digitised voltage and current signals into the frequency domain. This may be performed using a FFT.

The processing system may include processing electronics for performing the conversion.

15

The processing system is generally adapted to:

- a) Receive the converted voltage and current signals; and,
- b) Determine the impedance of the subject at each of the number of frequencies.

20 The processing system can be further adapted to determine the variation in the impedance with the frequency of the applied signal.

The processing system can be further adapted to generating a graphical representation of the variation in the impedance with the frequency of the applied signal.

25

In a third broad form the present invention provides a processing system for use in apparatus for determining the impedance of a subject, the processing system being adapted to:

- 30 a) Receive a digitised current signal representing the current flow through the subject at a number of frequencies for an applied electrical signal representing a range of superposed frequencies;
- b) Receive a digitised voltage signal representing the voltage across the subject at a number of frequencies within the range;

- c) Convert each digitised signal into the frequency domain; and,
- d) Determine the impedance of the subject at each of the number of frequencies.

The conversion can be performed using a FFT.

5

The processing system can include processing electronics for performing the conversion.

The processing system generally includes a processor for determining the impedance.

10 10 The processor can be further adapted to determine the variation in the impedance with the frequency of the applied signal.

The processing system typically includes a display, the processor being adapted to generating a graphical representation of the variation in the impedance with the frequency

15 15 of the applied signal.

In a fourth broad form the present invention provides a computer program product for determining the impedance of a subject, the computer program product including computer executable code which when executed by a suitable processing system causes the
20 processing system to operate as the processing system of the third broad form of the invention.

Brief Description of the Drawings

An example of the present invention will now be described with reference to the
25 accompanying drawings, in which: -

Figure 1 is a schematic diagram of an example of apparatus for implementing the present invention;

Figure 2 is a schematic diagram of the relationship between the Cartesian and Polar
30 impedance notation;

Figure 3 is a schematic diagram of an example of the processing system of Figure 1;

Figure 4 is a schematic diagram of a specific example of apparatus for implementing the invention; and,

Figure 5 is a schematic diagram of a specific example of a signal generator for use in the apparatus of Figures 1 or 5.

Detailed Description of the Preferred Embodiments

5 An example of apparatus suitable for performing the present invention is shown in Figure 1.

As shown the apparatus is formed from a signal generator 1, coupled to a body 2, such as a human subject, or the like, via electrodes 3, 4. A current detector 5 is coupled to the signal 10 generator 1 and one of the electrodes 3, with a voltage detector 6 being coupled to the body 2 via respective electrodes 6, 7, as shown. Signals from the current and voltage detectors 5, 7 are transferred to a processing system 9 for subsequent processing.

In use, the signal generator operates to apply an electrical signal to the body 2, via the 15 electrodes 3, 4. The current flow through the body is measured using the current detector 5, and transferred to the processing system 9. Simultaneously, the voltage generated across the body is measured using the voltage detector 6, and transferred to the processing system 9, thereby allowing the processing system 9 to determine the impedance of the body 2.

20

In particular, the impedance is calculated using the formula:

$$Z = V/I \quad (1)$$

Where: Z = impedance;

25 V = voltage; and,

I = current.

For complex impedance, each of these three values is represented by a complex vector. A complex vector can be represented in two ways, using either polar or Cartesian 30 coordinates. Polar notation uses the vector's length (Z) and it's phase (θ). The same information can also be described using Cartesian coordinates where the vector's X component is described as Resistance (R), and Y component is described as Reactance (X_c). This is shown for example in Figure 2.

The impedance of the body 2 can be measured at one particular frequency f by applying a pure sine wave current having the frequency f to the body and measuring the applied current and the voltage developed across the body 2. The determined voltage and current can then be used to determine the impedance.

5

If the calculations are to be performed digitally, the current and voltage measurements need to be sampled at a measurement rate of at least $2 \times f$, but realistically, to achieve good performance, a typical measurement rate should be higher, for example at $5 \times f$. This is required to prevent problems with aliasing of the sampled signals. It will be appreciated 10 that the more measurements taken, the more accurate the subsequent calculations will be. A typical number would be in the region of several thousand measurement points.

The apparatus described above operates to perform multiple frequency impedance measurements thereby allowing the system to determine the impedance for a number of 15 different applied frequencies of signal f_1, f_2, \dots, f_n . In order to achieve this, the apparatus uses the principle of superposition to allow the impedance calculations to be performed for multiple frequencies simultaneously.

In one example, this is achieved by having the signal generator generate an electrical signal 20 formed from the summation of multiple sine waves. Accordingly, the signal generator operates to superpose a number of sine waves and use these to form the electrical signal to be applied to the body 2. Ideally the resulting electrical signal should be formed from a superposition of a number of waves each having an equal amplitude.

25 The resulting current and voltages across the body 2 are then transferred to the processing system 9 to allow the processing system 9 to determine the impedance. Accordingly, it will be appreciated that any form of suitably adapted processing system may be used.

An example of suitable processing system is shown generally in Figure 3. In particular, 30 the processing system 9 includes a processor 10, a memory 11, an optional input/output (I/O) device 12, such as a keyboard and monitor, or the like, and an interface 13 coupled together via a bus 14. In use, the interface 13 is adapted to receive signals from the current and voltage detectors 5, 8. The processor 10 then executes applications software stored in

the memory 11, to process the received signals.

Accordingly, it will be appreciated that the processing system may be formed from any one of a number of forms of processing system, such as a suitably programmed PC, Lap-top, 5 hand held PC, palm-top or the like. Alternatively, the processing system 9 may be formed from specialised hardware, such as an electronic touch sensitive screen coupled to suitable processor and memory.

In any event, the processing system 9 operates to perform the impedance calculations by 10 converting time-domain sequences of voltage and current measurements obtained from the current and voltage detectors 5, 8 into frequency-domain data. This is typically (and most efficiently) performed using a FFT. A single pure sine wave of frequency f in the time domain will appear as a thin single peak at frequency f in the frequency domain (frequency spectrum), with the height of the peak being proportional to the amplitude of 15 the sine wave in the time domain. The FFT will also provide the phase (θ) of the sine wave, referenced to the start of the measurement period.

If the FFT operation is performed on both the voltage and current measurements, two peaks will result in the frequency spectrum, at the same frequency, but at differing heights 20 corresponding to the amplitudes of the voltage and current sine waves. If these two heights are divided by each other, the impedance is determined (as given by the formula (1) above, $Z = V/I$).

If the phases are subtracted from each other, the phase of the impedance vector is 25 determined. In this way, both values needed to define the impedance vector, namely its length (Z) and phase (θ), are determined. That is, the impedance vector is obtained by two simple FFT operations, one divide, and one subtraction.

As mentioned above, the system allows the impedance to be calculated for multiple 30 frequencies of interest simultaneously. Accordingly, the electrical signal applied to the body 2 is formed from a superposition of multiple sine waves. Subsequently a FFT is performed on the measured current and the voltage, and a division and a subtraction is

carried out for each point in the frequency spectrum. It will be appreciated that this process can be performed very rapidly, typically within a few milliseconds.

The more sine waves that are superposed, the more points will be determined in the 5 resulting frequency spectrum, and the more accurate the resulting plots. This principle can be maximised by applying a ‘white noise’ current to the body to be measured. An ideal white noise source contains equal amplitudes of all frequencies of interest. Accordingly, the use of an “ideal white noise” would allow the measurement of impedance at any number of frequencies simultaneously.

10

However, generating ideal white noise can be problematic, and accordingly, it is typical for the sample length of the white noise to be selected based on factors, such as the processing power available and the resolution required.

15 For example, if the white noise is selected to have a sample length of 1024 points, this will give 1024 separate points in the frequency domain, generating 1024 points on the resulting Cole-Cole plot.

However, in embodiments of the invention with sufficient processing power and storage 20 memory, larger numbers of points can be used, giving a very high resolution. Thus, the larger the sample length, the higher the resolution of the resulting plot. However, it will be appreciated that the use of more sample points requires a corresponding increase in the processing power required to process the measured voltage and current signals. Thus, it is typical to select a sample length based on the implementation and the circumstances in 25 which the invention is implemented, to thereby allowing the highest resolution to be determined based on the processing power available. This allows a wide range of

Furthermore, for a practical measurement of impedance, the white noise needs to be ‘band-limited’ where it will only contain frequency components up to a certain frequency f . The 30 A/D conversion sample rate must be at least $2 \times f$, to avoid “aliasing” errors in the processing, but realistically should be around at least $5 \times f$ for a practical device.

A specific example of apparatus suitable for performing the invention will now be described with reference to the Figure 4.

In this example, the apparatus uses specialised digital electronics to perform the 5 functionality outlined above with respect to Figure 1.

In particular, in this example, the signal generator is formed from a pseudo-random voltage generator 15, coupled to a current source 17, which is in turn coupled to the body 2 via electrodes 20.

10

Two further electrodes 21 are coupled to an A/D converter 25 to form the voltage detector 8, with the current detector being formed from a current sampler 23 and an associated A/D converter 26.

15 15 The A/D converters 25, 26 are then coupled to processing electronics shown by the dotted lines, which may be implemented either as respective digital electronics, the processing system 9, or a combination of the two. In this example, separate digital electronics and a processor 35 are used, as will be described below.

20 20 Operation of the system will now be described. In particular, the pseudo-random voltage generator 15, delivers an analogue command voltage 16 to the current source 17. The current source 17 is responsive to the received command voltage 16 to generate a pseudo-random "white noise" current 18, which is comprised of multiple frequencies, and which is applied to the two electrodes 20.

25

The two electrodes 21 are used to measure the voltage 22 generated across the body 2, with the voltage 22 being digitised by the A/D converter 25. In addition to this, the current sampler 23 samples the pseudo-random current 18 and the resulting signal 24 is digitised by the A/D converter 26.

30

In this example, the A/D converters 25, 26 will obtain measurements of the voltage 22 and the current signal 24 at a frequency that is at least five times greater than the maximum frequency of the applied current 18. It will therefore be appreciated that the sampling

frequency will be selected based on the preferred implementation. Thus, for example, the sampling frequency may be between 4MHz and 5MHz, although any suitable frequency may be used depending on the circumstances. Furthermore, as mentioned above a range of sampling lengths may be used, although in one example, the sample lengths can be 1024
5 bits.

As mentioned above, it will be appreciated that the greater the sample length, and sample rate the more accurate the process will be. However, the use of larger a sample length and/or rate will lead to a corresponding increase in the amount of data processing that will
10 be required. Accordingly, the use of 1024 bit samples, and a sampling rate of between 4MHz and 5MHz are illustrative only, but are particularly useful for providing good accuracy, without requiring undue processing. As processing systems and other digital electronics improve, it will be appreciated that higher sample lengths and rates will be achievable without effecting the time taken to obtain the readings.
15

The digital signal 27 resulting from the A/D conversion of the voltage 22 undergoes a FFT operation 30, which generates real and imaginary voltage components 33, 34 for multiple frequencies. Similarly the digital signal 28 output from the A/D conversion of the signal 24 undergoes a FFT operation 29, which generates real and imaginary components 31, 32
20 multiple frequencies.

It will be appreciated that the performance of the FFTs may be performed by the processor 10, or may alternatively be performed by separate processing electronics, as shown in this example. In any event, it will be appreciated that the signals received from the A/D
25 converters 25, 26 may need to be temporarily stored, for example in either the memory 11, or separate memory such as a shift register or the like, before being processed.

These real and imaginary components 31, 32, 33 and 34 generated by the FFT are transferred to the processor 10, where the resistive and reactive components 36, 37 of the
30 impedance for multiple frequencies are determined.

The resistive and reactive components 36, 37 can then be further processed and analysed in the processor 10 and a zero-infinity kHz parallel multiple frequency plot, also referred to as a Cole-Cole plot, can be generated before being displayed on the output device 12.

- 5 The resultant data shown generally at 39 can be transferred to the memory 11 for storage, or can be transferred to an external device 40, or the processor 10 for further processing and analysis. This includes the averaging of results, or the like, as will be described in more detail below.
- 10 In the preferred embodiment, the components 15, and 29-38 may be accomplished using digital circuitry, or suitably programmed processing systems.

An example of a circuit suitable for use as the pseudo-random voltage generator 15, used for the generation of band-limited white noise, will now be described with reference to 15 Figure 5.

In particular, the circuit includes a fixed frequency clock 41, a serial shift register 42, an XOR gate 45, a serial input - parallel output shift register 47, and a D/A converter 49.

- 20 In use, the fixed frequency clock 41 clocks the serial shift register 42. Several signals 44 from various points in the serial shift register 42 are fed into the XOR gate 45, the output 46 of which is fed back to the input of the serial shift register 42.

25 The output 43 from the serial shift register 42 is fed to the input of the serial input - parallel output shift register 47, which is also clocked from the clock 41. After the required number of bits appropriate for the correct operation of the D/A converter 49 have been shifted into the shift register 47 the parallel output 48 from the shift register 47 is sent to the D/A converter 49, which then generates the analogue command voltage 16.

- 30 Accordingly, the above described circuit shows a Linear Feedback Shift Register (LFSR) circuit that produces a pseudo-random digital sequence that is fed into the D/A converter 49. However, it will be appreciated that this represents only one technique for generating white noise, and other techniques can be used.

In the above example however, the random signal is based on a sequence that forms the contents of the serial shift register 42. This sequence should be of such a length that it does not repeat until after many successive measurements, which may be achieved for 5 example by providing a signal that is 100 bits in length, which will typically lead to a repeat time of several hundred million years.

This is important because one preferred embodiment is to use the apparatus described above to perform the measurement of the impedance over multiple frequencies a large 10 number of times and then average the result.

In particular, it will be appreciated that in the idealised case, the signal applied to the body 2 has an equal amplitude for each applied frequency. Accordingly, the relative magnitudes obtained for impedance measurements at different frequencies will not be influenced by 15 the applied signal.

However, if a random signal is used, it will be appreciated that the magnitude of the signal will vary from instant to instant. Accordingly, the impedance measured at any one time will depend to a degree on the magnitude of the applied signal at that time.

20

If the signal is truly random, then repeating impedance measurements a number of times and then averaging the results, will average out any variations in the resulting impedance that has arisen due to peaks and troughs in the applied signal.

25 However, in a pseudo random signal, it will be appreciated that any variations in the magnitude of the applied signal will repeat with some time period. Accordingly, if the sampling rate and repeat period happened to coincide, this may lead to exaggerated impedance measurements being obtained. By ensuring that the repeat time for the applied signal is significantly greater than the time periods over which measurements are taken, 30 this problem is avoided.

Accordingly, it will be appreciated that the above described systems can be used to determine the impedance of a body at several frequencies simultaneously. This vastly

reduces the length of time required to determine the impedance of an body, and in particular can reduce the time taken from several minutes achieved with existing techniques to a matter of milliseconds.

5 This in turn, allows repeat measurements to be performed over a short time period, such as a number of seconds, allowing the results from several readings to be averaged, thereby resulting in even more accurate results.

Another advantage of this invention is that the circuit required to undertake the above
10 operations can be almost entirely digital, giving the usual advantages of digital circuitry, namely, repeatability, reliability, no drift over either temperature or time, and simplicity of operation.

Persons skilled in the art will appreciate that numerous variations and modifications will
15 become apparent. All such variations and modifications which become apparent to persons skilled in the art, should be considered to fall within the spirit and scope that the invention broadly appearing before described.

Thus, for example, it will be appreciated that the above described techniques may be
20 utilised to determine the bioelectric impedance of an biological sample, and is not restricted to applications for humans, or the like.

THE CLAIMS DEFINING THE INVENTION ARE AS FOLLOWS:

- 1) A method of determining the impedance of a subject, the method including:
 - a) Applying an electrical signal representing a range of superposed frequencies;
 - b) Determining for a number of frequencies within the range:
 - 5 i) The current flow through the subject; and,
 - ii) The voltage across the subject; and,
 - c) Determining the impedance of the subject at each of the number of frequencies.
- 2) A method according to claim 1, the method including:
 - a) Generating component signals, each component signal having a respective one of
10 the number of frequencies; and,
 - b) Superposing the component signals to generate the electrical signal.
- 3) A method according to claim 1, the electrical signal being formed from white noise.
- 4) A method according to claim 3, the method including:
 - a) Generating the white noise using a Linear Feedback Shift Register (LFSR) circuit
15 to produce a pseudo-random digital sequence; and,
 - b) Converting the pseudo-random digital sequence to an analogue signal using a digital to analogue (D/A) converter; and,
 - c) Applying the analogue signal to the subject.
- 5) A method according to any one of the claims 1 to 4, the method of determining the
20 current flow including:
 - a) Sampling the current of the electrical signal applied to the subject; and,
 - b) Converting the current signal to a digitised current signal.
- 6) A method according to claim 5, the method of determining the voltage including:
 - a) Obtaining a signal representing the voltage generated across the subject;
 - 25 b) Converting the voltage signal to a digitised voltage signal.
- 7) A method according to claim 6, the method including digitising the current and voltage signals by sampling the signals at a predetermined sample rate.
- 8) A method according to claim 6 or claim 7, the method including digitising the current and voltage signals by sampling the signals with a predetermined sample length.
- 30 9) A method according to any one of the claims 6 to 8, the method including converting each of the digitised voltage and current signals into the frequency domain.
- 10) A method according to claim 9, the conversion being performed using a Fast Fourier Transform (FFT).

- 11) A method according to claim 9 or claim 10, the method including using a processing system to:
 - a) Receive the converted voltage and current signals; and,
 - b) Determine the impedance of the subject at each of the number of frequencies.
- 5 12) A method according to claim 11, the processing system being further adapted to determine the variation in the impedance with the frequency of the applied signal.
- 13) A method according to claim 12, the method further including generating a graphical representation of the variation in the impedance with the frequency of the applied signal.
- 10 14) A method of determining the impedance of a subject, the method being substantially as hereinbefore described.
- 15) Apparatus for determining the impedance of a subject, the apparatus including:
 - a) A signal generator for applying an electrical signal representing a range of superposed frequencies;
 - b) A voltage detector for determining the voltage across the subject at a number of frequencies within the range;
 - c) A current detector for determining the current flow through the subject at a number of frequencies within the range; and,
 - d) A processing system for determining the impedance of the subject at each of the number of frequencies.
- 20 16) Apparatus according to claim 15, the signal generator being adapted to:
 - a) Generate component signals, each component signal having a respective one of the number of frequencies; and,
 - b) Superpose the component signals to generate the electrical signal.
- 25 17) Apparatus according to claim 15, the electrical signal being formed from white noise.
- 18) Apparatus according to claim 17, the signal generator including:
 - a) A shift register circuit to produce a pseudo-random digital sequence; and,
 - b) A D/A converter for converting the pseudo-random digital sequence to an analogue signal.
- 30 19) Apparatus according to claim 18, the signal generator including:
 - a) A shift register having an output coupled to the D/A converter; and,
 - b) An exclusive OR (XOR) gate adapted to:
 - i) Receive inputs from a number of predetermined locations in the first register;

- ii) Logically combine the inputs to generate an XOR output; and,
- iii) Provide the XOR output to an input of the shift register;

20) Apparatus according to claim 19, the signal generator including a second shift register, the second shift register being adapted to couple an output of the first shift register to an input of the D/A converter.

5 21) Apparatus according to any one of the claims 15 to 20, the current detector including:

- a) A current sampler coupled to the signal generator for sampling the current flowing through the subject; and,
- b) A current analogue to digital (A/D) converter for converting the sampled current to a digitised current signal.

10 22) Apparatus according to claim 21, the voltage detector including a voltage A/D converter coupled to the subject via a respective set of electrodes, the voltage A/D converter being adapted to generate a digitised voltage signal.

23) Apparatus according to claim 22, the current and voltage A/D converters being adapted to digitise the current and voltage signals by sampling the signals at a predetermined sample rate.

15 24) Apparatus according to claim 22 or claim 23, the current and voltage D/A converters being adapted to digitise the current and voltage signals by sampling the signals with a predetermined sample length.

20 25) Apparatus according to any one of the claims 22 to 24, the processing system being adapted to convert each of the digitised voltage and current signals into the frequency domain.

26) Apparatus according to claim 25, the conversion being performed using a FFT.

27) Apparatus according to claim 25 or claim 26, the processing system including 25 processing electronics for performing the conversion.

28) Apparatus according to any one of the claims 25 to 27, the processing system being adapted to:

- a) Receive the converted voltage and current signals; and,
- b) Determine the impedance of the subject at each of the number of frequencies.

30 29) Apparatus according to any one of the claims 15 to 28, the processing system being further adapted to determine the variation in the impedance with the frequency of the applied signal.

30) Apparatus according to any one of the claims 15 to 29, the processing system being further adapted to generating a graphical representation of the variation in the impedance with the frequency of the applied signal.

31) Apparatus for determining the impedance of a subject, the apparatus being substantially as hereinbefore described.

5 32) A processing system for use in apparatus for determining the impedance of a subject, the processing system being adapted to:

- a) Receive a digitised current signal representing the current flow through the subject at a number of frequencies, for an applied electrical signal representing a range of superposed frequencies;
- 10 b) Receive a digitised voltage signal representing the voltage across the subject at a number of frequencies within the range;
- c) Convert each digitised signals into the frequency domain; and,
- d) Determine the impedance of the subject at each of the number of frequencies.

15 33) A processing system according to claim 32, the conversion being performed using a FFT.

34) A processing system according to claim 32 or claim 33, the processing system including processing electronics for performing the conversion.

20 35) A processing system according to any one of the claims 15 to 28, the processing system including a processor for determining the impedance.

36) A processing system according to claim 35, the processor being further adapted to determine the variation in the impedance with the frequency of the applied signal.

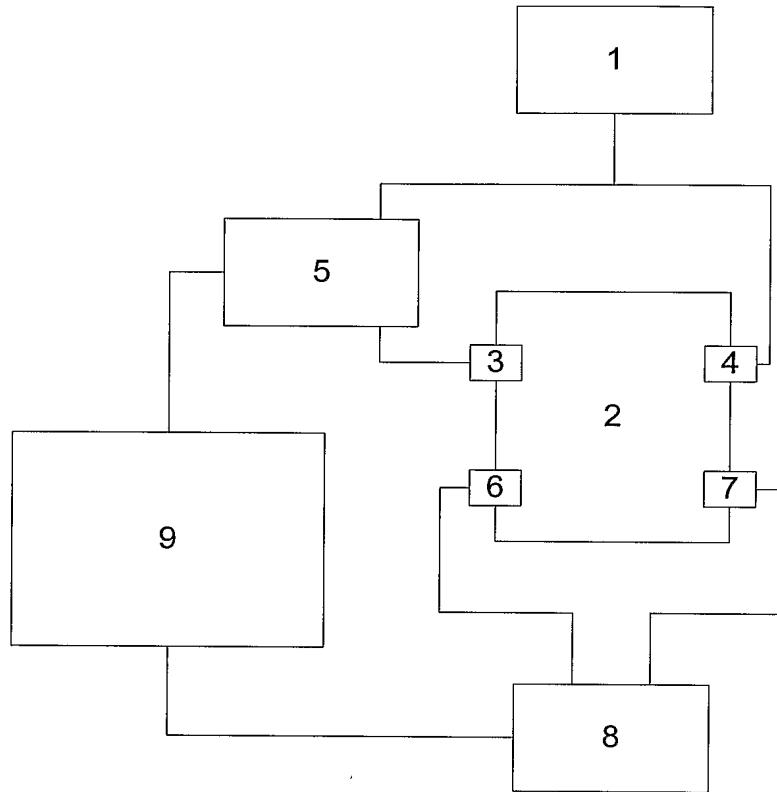
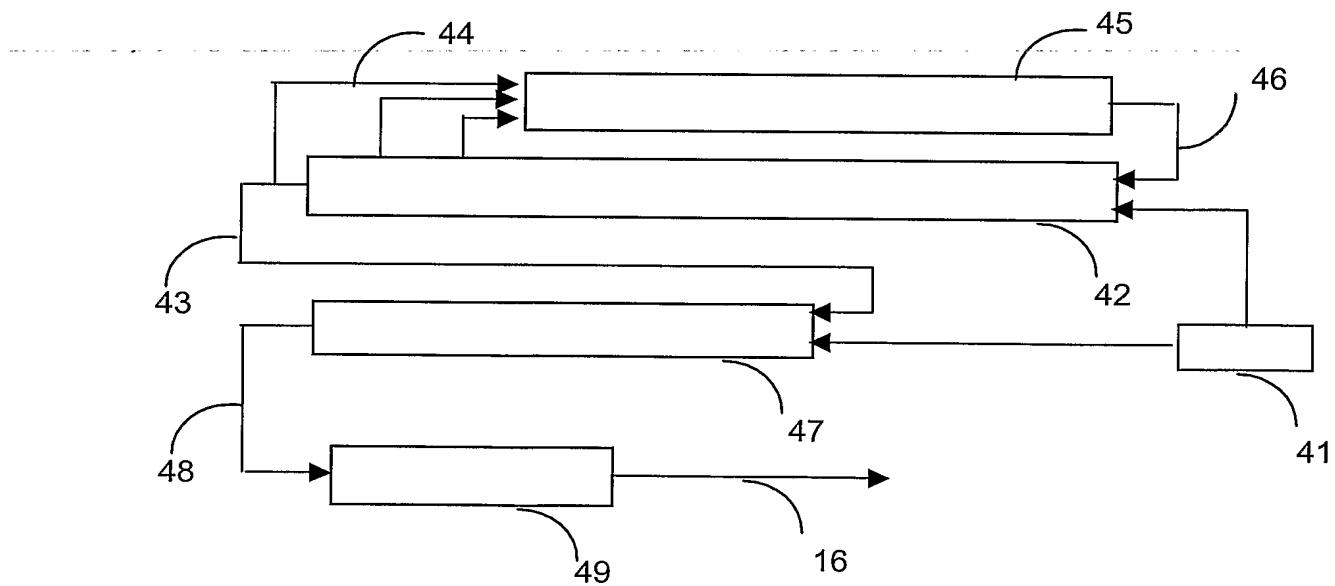
25 37) A processing system according to claim 36, the processing system including a display, the processor being adapted to generating a graphical representation of the variation in the impedance with the frequency of the applied signal.

38) A processing system for determining the impedance of a subject, the processing system being substantially as hereinbefore described.

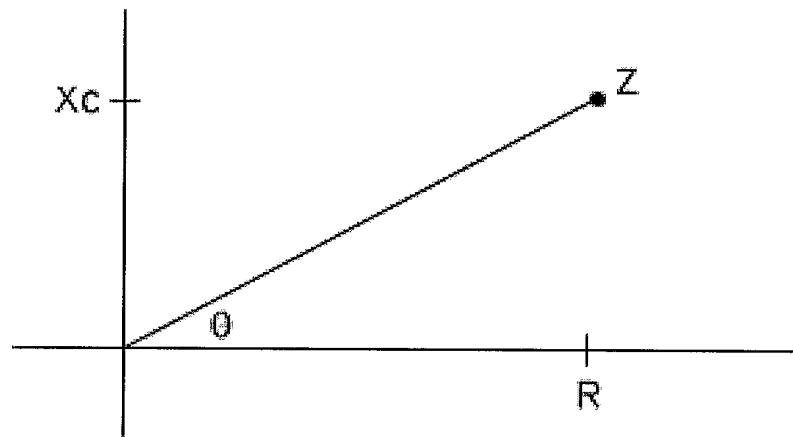
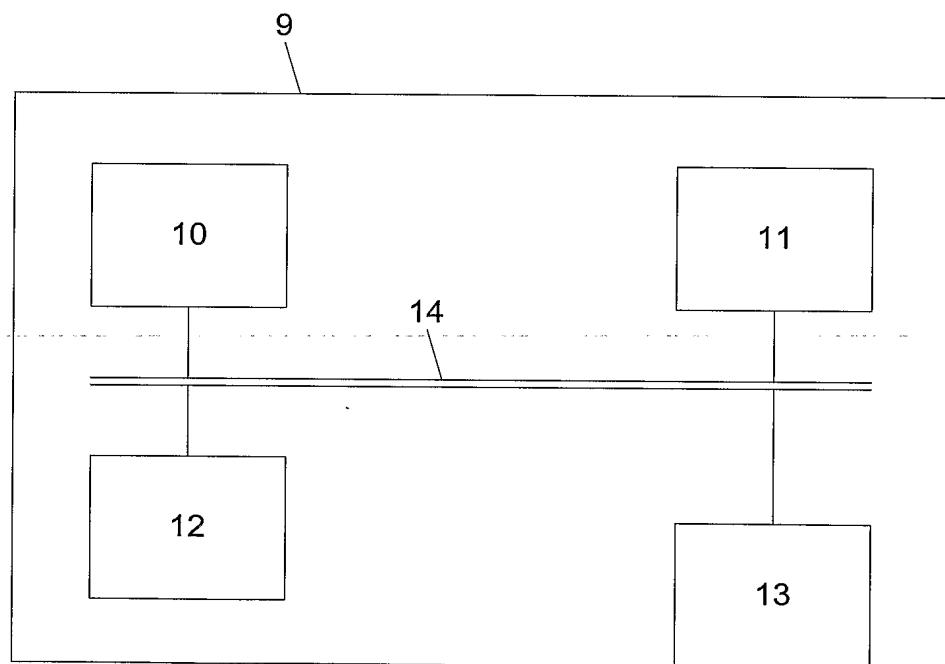
39) A computer program product for determining the impedance of a subject, the computer program product including computer executable code which when executed by a suitable processing system causes the processing system to operate as the processing system of any one of the claims 32 to 38.

30

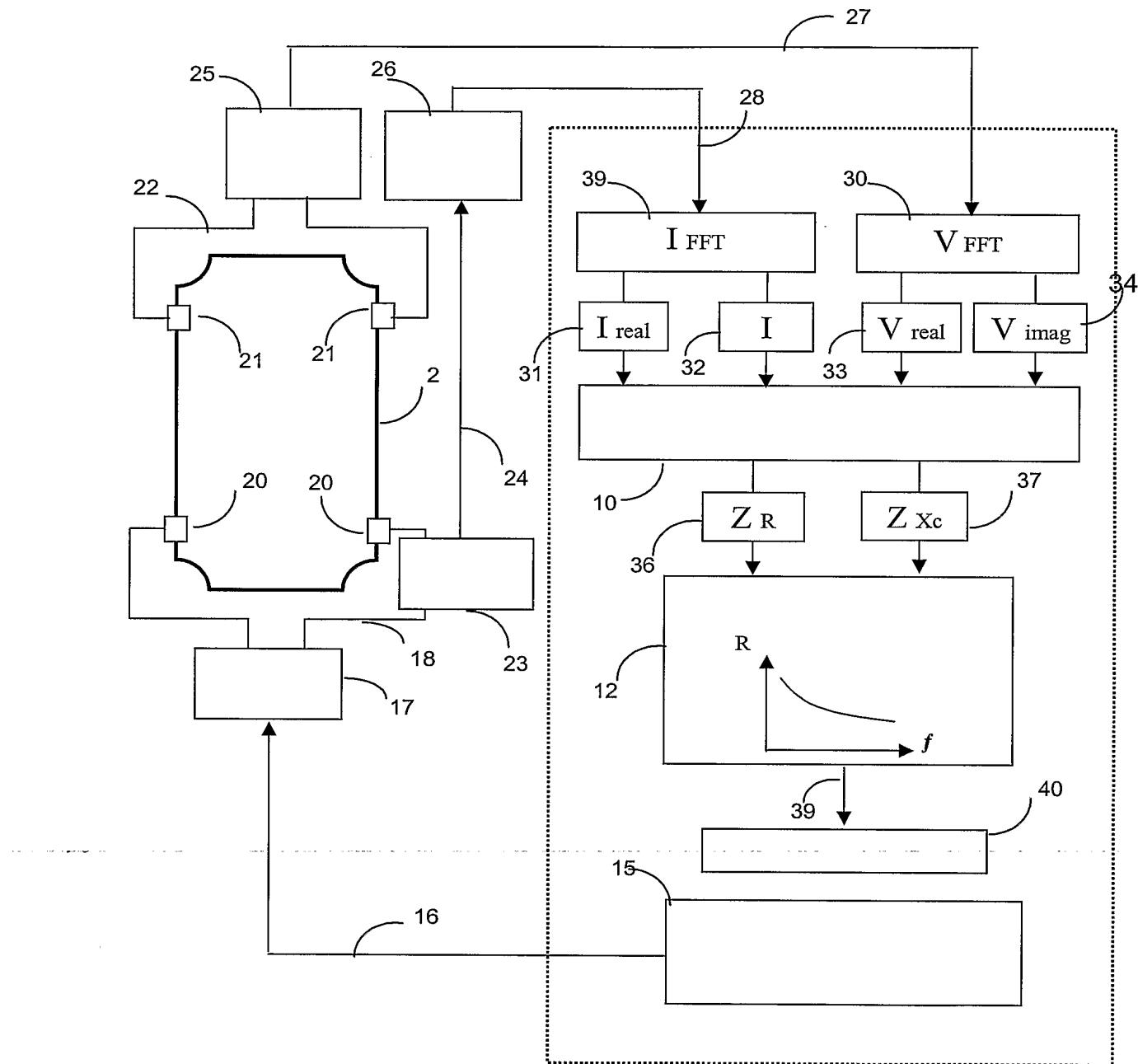
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**Fig. 1****Fig. 5**

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**Fig. 2****Fig. 3**

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**Fig. 4**

INTERNATIONAL SEARCH REPORT

International application No.
PCT/AU2003/001566

A. CLASSIFICATION OF SUBJECT MATTER																						
Int. Cl. ⁷ : A61B 5/05																						
According to International Patent Classification (IPC) or to both national classification and IPC																						
B. FIELDS SEARCHED																						
Minimum documentation searched (classification system followed by classification symbols) SEE ELECTRONIC DATABASES CONSULTED																						
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched																						
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) DWPI JAPIO MEDLINE: bioimpedance bia mfbia bioelectric impedance reactance inductance admittance phase triple multi sweep scan spectrum plural frequency simultaneous superimpose same A61B A61N G10K G01N																						
C. DOCUMENTS CONSIDERED TO BE RELEVANT																						
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.																				
X	JP 09-051884 A (SEKISUI CHEMICAL CO LTD) 25 February 1997 abstract	1-39																				
X	Thomas BJ, Future Technologies. Asia Pacific Journal Clinical Nutr, (1995), 4: 157-159. Page 158	1, 2, 14-16, 31-39																				
X	Ellis KJ et al, Human hydrometry: comparison of multifrequency bioelectrical impedance with ² H ₂ O and bromine dilution. Journal of Applied Physiology, (1998), 85(3): 1056-1062. Page 1057	1, 2, 14-16, 31-39																				
<input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C		<input checked="" type="checkbox"/> See patent family annex																				
<p>* Special categories of cited documents:</p> <table style="width: 100%; border-collapse: collapse;"> <tr> <td style="width: 15%;">"A"</td> <td>document defining the general state of the art which is not considered to be of particular relevance</td> <td style="width: 15%;">"T"</td> <td>later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</td> </tr> <tr> <td>"E"</td> <td>earlier application or patent but published on or after the international filing date</td> <td>"X"</td> <td>document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</td> </tr> <tr> <td>"L"</td> <td>document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</td> <td>"Y"</td> <td>document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art</td> </tr> <tr> <td>"O"</td> <td>document referring to an oral disclosure, use, exhibition or other means</td> <td>"&"</td> <td>document member of the same patent family</td> </tr> <tr> <td>"P"</td> <td>document published prior to the international filing date but later than the priority date claimed</td> <td></td> <td></td> </tr> </table>			"A"	document defining the general state of the art which is not considered to be of particular relevance	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention	"E"	earlier application or patent but published on or after the international filing date	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone	"L"	document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art	"O"	document referring to an oral disclosure, use, exhibition or other means	"&"	document member of the same patent family	"P"	document published prior to the international filing date but later than the priority date claimed		
"A"	document defining the general state of the art which is not considered to be of particular relevance	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention																			
"E"	earlier application or patent but published on or after the international filing date	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone																			
"L"	document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art																			
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"P"	document published prior to the international filing date but later than the priority date claimed																					
Date of the actual completion of the international search 27 January 2004	Date of mailing of the international search report 10 FEB 2004																					
Name and mailing address of the ISA/AU AUSTRALIAN PATENT OFFICE PO BOX 200, WODEN ACT 2606, AUSTRALIA E-mail address: pct@ipaaustralia.gov.au Facsimile No. (02) 6285 3929	Authorized officer  MATTHEW FORWARD Telephone No : (02) 6283 2606																					

INTERNATIONAL SEARCH REPORT

International application No.

PCT/AU2003/001566

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	Jones CH et al, Extracellular fluid volume determined by bioelectric impedance and serum albumin in CAPD patients. <i>Nephrology Dialysis Transplantation</i> (1998), 13: 393-397. Page 393	1, 2, 14-16, 31-39
X	Woodrow G et al, Effects of icodextrin in automated peritoneal dialysis on blood pressure and bioelectrical impedance analysis. <i>Nephrology Dialysis Transplantation</i> (2000), 15: 862-866. Page 863	1, 2 14-16, 31-39
A	US 5280429 A (WITHERS) 18 January 1994	
A	EP 0865763 A2 (NTE, S.A) 23 September 1998	
A	JP 2000-139867 A (SEKISUI CHEMICAL CO. LTD.) 23 May 2000 Abstract	
A	WO 2001/067098 A1 (BTG INTERNATIONAL LIMITED) 13 September 2001	
P, A	US 6532384 B1 (FUKUDA) 11 March 2003	

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No.

PCT/AU2003/001566

This Annex lists the known "A" publication level patent family members relating to the patent documents cited in the above-mentioned international search report. The Australian Patent Office is in no way liable for these particulars which are merely given for the purpose of information.

Patent Document Cited in Search Report				Patent Family Member			
JP	09-051884	NO	FAMILY				
US	5280429	CA	2102231	EP	0587646	JP	05-280429
		WO	1992/019153				
EP	0865763	ES	2151774	US	6151523		
JP	2000-139867	NO	FAMILY				
WO	2001/067098	AU	37555/01	CA	2401508	EP	1259806
		US	2003/0105411				
US	6532384	CN	1287823	EP	1080686	JP	2000-070273
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